

The listing of claims will replace all prior versions and listing of claims in the application:

**Listing of Claims:**

Claim 1. (Withdrawn) A method for treating a patient, who is to receive a cytotoxic agent, which comprises the steps of:

- (a) determining the circulating suramin concentration in said patient;
- (b) administering suramin, if required, in a required dose to establish a low circulating concentration of suramin in said patient of below about 200  $\mu$ M; and
- (c) administering said cytotoxic agent to said patient when said low circulating concentration of suramin of below about 200  $\mu$ M is present in said patient.

Claim 2. (Withdrawn) The method of claim 1, wherein said low dose of circulating suramin is between about 10 and 200  $\mu$ M.

Claim 3. (Withdrawn) The method of claim 2, wherein said low dose of circulating suramin is between about 10 and 50  $\mu$ M.

Claim 4. (Withdrawn) The method of claim 1, wherein the required dose of suramin can be determined in step (b) by the steps of:

- (b1) determining the gender and the squared value of the body surface area (BSA) of said patient;
- (b2) determining the time elapsed, in days, since the initiation of the last suramin treatment; and
- (b3) calculating the dose of low dose suramin using a nomogram that shows the dose according to the parameters of gender, squared value of body surface, and elapsed days since last suramin treatment.

Claim 5. (Withdrawn) The method of claim 4, wherein said nomogram comprises:

Nomogram For Calculating Suramin Dose		
	FACTOR	
	Man	Woman
Cycle 1*	125	125
Days since the administration of the first dose of previous cycle		

	FACTOR	
7	39	33
8	43	37
9	47	40
10	51	44
11	55	47
12	58	50
13	61	53
14	64	56
15	67	58
16	70	61
17	72	63
18	74	66
19	77	68
20	79	70
21	80	72
22	82	74
23	84	75
24	86	77
25	87	79
26	88	80
27	90	82
28	91	83
29	92	84
30	93	86
31	94	87
32	95	88
33	96	89
34	97	90
35	98	91
36	98	92
37	99	93
38	100	94

39	100	95
41	102	96
42	102	97
44	103	98
47	104	100
49	105	101
52	106	102
55	106	103

where:

$$\text{First cycle dose (mg)} = \frac{(21.4 * 5.13 * \text{BSA}^2)}{e^{-(0.0026 \text{ or } 0.0022 * 48)}} = \text{FACTOR} * \text{BSA}^2 \quad \text{Eq. 15}$$

and

$$\text{Subsequent cycle dose} = \text{First dose} * (1 - e^{-k^* t}) = 125 * \text{BSA}^{2*} (1 - e^{-k^* t}) \quad \text{Eq. 16.}$$

Claim 6. (Withdrawn) The method of claim 1, wherein said cytotoxic agent is one or more of an anti-microtubule agent, a topoisomerase I inhibitor, a topoisomerase II inhibitor, an anti-metabolite, a mitotic inhibitor, an alkylating agent, an intercalating agent, an agent capable of interfering with a signal transduction pathway, an agent that promotes one or more of apoptosis or necrosis, an interferon, an interleukin, a tumor necrosis factor, or radiation.

Claim 7. (Withdrawn) The method of claim 6, wherein said cytotoxic agent is one or more of paclitaxel, vincristine, vinblastine, vindesine, vinorelbine, docetaxel, topotecan, camptothecin, irinotecan hydrochloride, doxorubicin, etoposide, mitoxantrone, daunorubicin, idarubicin, teniposide, amsacrine, epirubicin, merbarone, piroxantrone hydrochloride, 5-fluorouracil, methotrexate, 6-mercaptopurine, 6-thioguanine, fludarabine phosphate, cytosine arabinoside, trimetrexate, gemcitabine, acivicin, alanosine, pyrazofurin, N-Phosphoracetyl-L-Aspartate (PALA), pentostatin, 5-azacitidine, 5-Aza-2'-deoxycytidine, adenosine arabinoside, cladribine, fltoraful, UFT (combination of uracil and fltoraful), 5-fluoro-2'-deoxyuridine, 5'-deoxy-5-fluorouridine, tiazofurin, Xeloda (Capecitabine), cisplatin, carboplatin, oxaliplatin, mitomycin C, BCNU, melphalan, thiotepa, busulfan, chlorambucil, plicamycin, dacarbazine, ifosfamide phosphate,

cyclophosphamide, nitrogen mustard, uracil mustard, pipobroman, 4-ipomeanol, dihydronerperone, spiromustine, geldanamycins, cytochalasins, depsipeptide, leuprolide (e.g., Lupron), ketoconazole, tamoxifen, goserelin, flutamide, 4'-cyano-3-(4-fluorophenylsulphonyl)-2-hydroxy-2-methyl-3'-(trifluoromethyl) propionanilide, Herceptin, anti-CD20 (Rituxan), C225, Iressa, interferon alpha, interferon beta, interferon gamma, interleukin 2, interleukin 4, interleukin 12, tumor necrosis factors, radiation, hydroxyurea, azathioprine, aminopterin, trimethoprin, pyrimethamine, pyritrexim, DDMP (2,4 diamino-5(3',4' dichlorophenyl)6 methylpyrimidine), 5,10-dideazatetrahydrofolate, 10-propargyl-5,8 dideazafolate (CB3717), 10-ethyl-10-deaza-aminopterin, deoxycytidine, 5-aza-cytosine arabinoside, N-4-palmitoyl-ara C, 2'-azido-2'-deoxy-ara C, N4-behenoyl-ara C, CCNU (lomustine), estramustine, MeCCNU, triethylene melamine, trenimon, dimethyl busulfan, streptozotocin, chlorozotocin, procarbazine, hexamethylmelamine (Altretamine), pentamethylmelamine (PMM), tetraplatin, oxaliplatin, platinum-DACH, aziridinylbenzoquinone (AZQ), bleomycin, tallysomycin S<sub>10</sub><sup>b</sup>, liblomycin, pepleomycin, asparaginase (Elspar), pegaspargase (Oncaspar), Cladrabine (leustatin), porfimer sodium (Photofrin), amonofide, deoxyspergualin, dihydronerperone, flavone acetic acid, gallium nitrate, or hexamethylene bisacetamine (HMBA).

Claim 8. (Withdrawn) The method of claim 1, wherein a suramin dose is administered such that a concentration of between about 10 to about 50  $\mu$ M over 48 hours is achieved in a patient.

Claim 9. (Withdrawn) The method of claim 1, wherein the patient is a mammal.

Claim 10. (Withdrawn) The method of claim 9, wherein the patient is a human.

Claim 11. (Withdrawn) The method of claim 1, wherein the patient has a tumor.

Claim 12. (Withdrawn) The method of claim 7, wherein the cytotoxic agent is one or more of carboplatin or paclitaxel.

Claim 13. (Withdrawn) The method of claim 1, wherein two-thirds of the therapeutically effective amount of suramin is given on the first day and the remaining one-third of the therapeutically effective amount of suramin is given about 24 hours later.

Claim 14. (Withdrawn) The method of claim 1, wherein the required dose of suramin can be determined in step (b) by the steps of:

- (b1) determining the squared value of the body surface area (BSA) of said patient;
- (b2) determining the time elapsed, in days, since the initiation of the last suramin treatment; and
- (b3) calculating the dose of low dose suramin using a nomogram that shows the dose according to the parameters of gender, squared value of body surface, and elapsed days since last suramin treatment.

Claim 15. (Withdrawn) The method of claim 14, wherein said nomogram comprises:

Nomogram For Calculating Suramin Dose

Cycle 1*	125
Days since the administration of the first dose of previous cycle	FACTOR
7	39
8	43
9	47
10	51
11	55
12	58
13	61
14	64
15	67
16	69
17	72
18	74

19	76
20	78
21	80
22	82
23	84
24	86
25	87
26	88
27	90
28	91
29	92
30	93
31	94
32	95
33	96
34	97
35	98
36	98
37	99
38	100
39	100
41	102
42	102
44	103
47	104

49	105
52	106
55	106

where:

$$\text{First cycle dose (mg)} = \frac{(21.4 * 5.13 * \text{BSA}^2)}{e^{-(0.0026 \text{ or } 0.0022 * 48)}} = \text{FACTOR} * \text{BSA}^2 \quad \text{Eq. 15}$$

and

$$\text{Subsequent cycle dose} = \text{First dose} * (1 - e^{-k^* t}) = 125 * \text{BSA}^2 * (1 - e^{-k^* t}) \quad \text{Eq. 16.}$$

Cancel claims 16-21, inclusive.

Claim 22. (Original) A kit for carrying out the combined administration of suramin with one or more cytotoxic agents, comprising:

- (a) suramin formulated in a pharmaceutical carrier; and
- (b) instructions for therapeutic use of said suramin in combination with said cytotoxic agent(s) in one or more of inhibiting growth, proliferation of tumor cells, or inducing killing of tumor cells, calling for:
  - (i) administering suramin, if required, in a required dose to establish a low circulating concentration of suramin in said patient of below about 200  $\mu\text{M}$ ; and
  - (ii) administering said chemotherapeutic agent to said patient when said low circulating concentration of suramin of below about 200  $\mu\text{M}$  is present in said patient.

Claim 23. (Original) The kit of claim 22, wherein said instructions include a method for determining a therapeutically effective amount of suramin.

Claim 24. (Original) The kit of claim 23, wherein the instructions for determining a therapeutically effective amount of suramin comprise a nomogram.

Claim 25. (Canceled)

Claim 26. (Original) The kit of claim 22, wherein one of the cytotoxic agents is carboplatin.

Claim 27. (New) The kit of claim 22, wherein said low circulating concentration of suramin is between about 10 and 200  $\mu$ M.

Claim 28. (New) The kit of claim 27, wherein said low dose of circulating suramin is between about 10 and 50  $\mu$ M.

Claim 29. (New) The kit of claim 22, wherein the required dose of suramin can be determined in step (b) by the steps of:

- (b1) determining the squared value of the body surface area (BSA) of said patient;
- (b2) determining the time elapsed, in days, since the initiation of the last suramin treatment; and
- (b3) calculating the dose of low dose suramin using a nomogram that shows the dose according to the parameters of squared value of body surface, and elapsed days since last suramin treatment.

Claim 30. (New) The kit of claim 29, wherein said nomogram comprises:

Nomogram For Calculating Suramin Dose

Cycle 1*	125
Days since the administration of the first dose of previous cycle	FACTOR
7	39
8	43
9	47
10	51
11	55
12	58
13	61

14	64
15	67
16	69
17	72
18	74
19	76
20	78
21	80
22	82
23	84
24	86
25	87
26	88
27	90
28	91
29	92
30	93
31	94
32	95
33	96
34	97
35	98
36	98
37	99
38	100

39	100
41	102
42	102
44	103
47	104
49	105
52	106
55	106

where:

$$\text{First cycle dose (mg)} = \frac{(21.4 * 5.13 * \text{BSA}^2)}{e^{-(0.0026 \text{ or } 0.0022 * 48)}} = \text{FACTOR} * \text{BSA}^2 \quad \text{Eq. 15}$$

and

$$\text{Subsequent cycle dose} = \text{First dose} * (1 - e^{-k^* t}) = 125 * \text{BSA}^{2*} (1 - e^{-k^* t}) \quad \text{Eq. 16.}$$

Claim 31. (New) The kit of claim 22, wherein said cytotoxic agent is one or more of an anti-microtubule agent, a topoisomerase I inhibitor, a topoisomerase II inhibitor, an anti-metabolite, a mitotic inhibitor, an alkylating agent, an intercalating agent, an agent capable of interfering with a signal transduction pathway, an agent that promotes one or more of apoptosis or necrosis, an interferon, an interleukin, a tumor necrosis factor, or radiation.

Claim 32. (New) The kit of claim 22, wherein a suramin dose is administered such that a concentration of between about 10 to about 50  $\mu\text{M}$  over 48 hours is achieved in a patient.

Claim 33. (New) The kit of claim 22, wherein two-thirds of the therapeutically effective amount of suramin is given on the first day and the remaining one-third of the therapeutically effective amount of suramin is given about 24 hours later.